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# The economic burden of prostate cancer in Eswatini

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# **Abstract**

**Background:** Prostate cancer is the fifth cause of cancer mortality among men worldwide. However, there is limited data on costs associated with prostate cancer in low- and middle-income countries particularly in the sub-Saharan region. From a societal perspective, this study aims to estimate the cost of prostate cancer in Eswatini.

**Methods:** This prevalence–based cost-of-illness study used diagnosis specific data from national registries to estimate costs associated to prostate cancer during 2018. The prevalence-based approach was used employing both top down and bottom up costing approaches. Costs data included health care utilization, transport, sick leave days and premature death.

**Results:** The total annual cost of prostate cancer was \$6.2 million (ranging between \$4.7 million and 7.8 million estimated with lower and upper bounds). Average cost-per patient for radiotherapy, chemotherapy and other non-medical direct costs (transport and lodging) were the highest cost drivers recording \$16,648, \$7,498 and \$5,959 respectively whilst indirect costs including productive loss due to sick leave and pre-mature mortality was estimated at \$58,320 and \$113,760 respectively. Cost of managing prostate cancer increased with advanced disease and costs were highest for prostate cancer stages III and IV recording \$1.1 million, \$1.9 million respectively.

**Conclusions:** Prostate cancer is a public health concern in Eswatini, and it imposes significant economic burden to the society. This finding point areas for policy makers to perform cost containment regarding therapeutic procedures for prostate cancer and the need for strategies to increase efficiencies in the health care systems for increased value for health care services.

**Keywords:** Prostate cancer, Cost-of-illness, Eswatini, Premature mortality, Prostate antigen test

# **Background**

Among cancers, prostate cancer is the third commonest cancer after breast and lung cancer and the fifth cause of cancer mortality among men [1, 2]. In 2018, the number of new cases increased from 1.1 million in 2012 to 1.3 million in 2018 accounting for about 7.1% of the total cancer cases globally and 15% among men [2]. The causes of prostate cancer is attributable to genetic

and environmental factors [2]. However, the incidence and mortality rate vary substantially within and across regions. Notably, high-income countries (HICs) reports high incidence rate compared to low- and -middle income countries (LMICs) [2]. In contrast, mortality rate is higher in developing countries particularly in sub-Saharan Africa regions [3]. The inequalities observed across regions with respect to prostate cancer incidence and mortality are in part linked to availability of effective screening and improved treatment modalities which are directly linked to resources availability [3, 4]. In Eswatini, compared to other common cancers, prostate cancer is ranked third accounting for 7.6% of total new cases 1074 in 2018 [5].

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Prostate cancer causes clinical and economic burden to patients and governments. Screening tests include prostate-specific antigen (PSA) and digital rectal examination (DRG) [6, 7]. A positive screening tests results indicate further investigation [6]. Whilst PSA is the frequent screening test, it has been argued that PSA could potentially cause harm by over diagnosing low risk cancers that otherwise would have remained without clinical consequences for life time if left untreated [8]. In turn, this increases costs for prostate cancer [9]. In Sweden, annual costs associated with prostate cancer (screening, diagnosis and treatment) was estimated at €281 million [9]. In Ontario, the mean per patient cost for prostate cancer-related medication was \$1211 [10]. In Iran, the total annual cost of prostate cancer was estimated at \$2900 million [11]. Other studies estimated the economic burden of prostate cancer along with other cancer type. A study focusing on European countries, ranked prostate cancer the fourth cancer disease to cause health care costs compared to lung (€18.8billion), breast cancer (€15 billion), colorectal cancer (€13.1 billion) [12]. Similarly, in Korea, prostate cancer was among the top four cancers attributing to economic burden of disease [13].

There is limited evidence on the economic burden of prostate cancer from LMICs. Estimation of the economic burden of disease provide insight on treatment modalities and associated costs. The study aims to investigate the societal cost of prostate cancer in Eswatini during 2018.

# Materials and methods

# Study area

Eswatini formerly known as Swaziland is a country in Southern African bordering South Africa and Mozambique with an estimated population of 1.2 million [14]. The country's economy is tied to South Africa and Eswatini's domestic currency (Lilangeni=SZL) is pegged at parity with South African currency (Rand=ZAR) such that Eswatini cannot conduct its own monetary policy [15]. Eswatini fiscal revenue largely depend on Southern African Customs Union (SACU) revenues and remittance flowing mainly from South Africa [16, 17]. SACU receipts account for about a third of Eswatini's total revenue and grants. However, over the past decades, SACU revenues have consistently declined leaving Eswatini's economy constrained. The country records high national level poverty rate and income inequality which does not commensurate with its middle-income status. The national poverty rate is 58.9% percent at the international \$1.90 poverty line and Gini index- a measure of inequality is 49.3 [17]. Eswatini ranks near the bottom of the World Bank's Human Capital Index, with a score of 0.37 in 2020. Eswatini health spending as a share of the total budget is estimated at 10.1% and health per capita is estimated at \$ 248 per annum [16]. Whilst Eswatini's health expenditure is comparatively higher to some other countries in the Southern African region, the country's health outcomes do not reflect its spending levels on health and its middle-income status. The health care service delivery is made up of public and private health care. Compared to the public, the private health care systems is better equipped both infrastructural and human resources however, at high health care costs. As such, private health care is accessed by less than 10% of the population, mainly those who owns health insurance [18].

Diagnostic and treatment capacity of conditions including cancer remains limited in the country mostly in the public health system. Through a government funded scheme namely Phalala, the Eswatini citizens are supported to access specialized health care services from neighboring countries mainly South Africa.

# Methods of costing

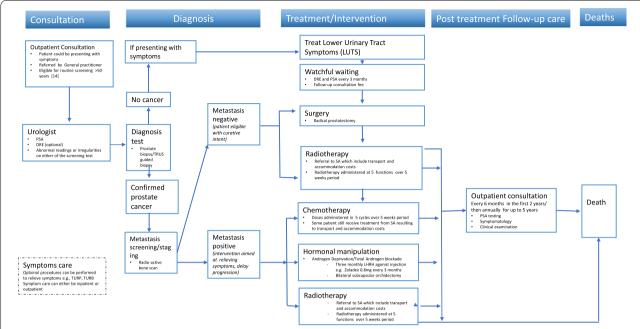
This is a Cost of Illness (CoI) study investigating costs of prostate cancer from the societal perspective [19]. CoI studies estimate disease specific costs [20]. The prevalence based approach, was used employing both top down and bottom up costing approaches [19, 21]. The cost estimation involved identification, quantification and valuation of resources used. The total costs for prostate cancer was calculated by multiplying identified resources quantities and the respective unit costs. All costs were presented in US\$ adjusted for 2018 (\$1= SZL14.5).

# Study population

Data on prostate cancer prevalence and mortality in 2018 was obtained from the National cancer registry [14]. The National Cancer Control Unit is led by the Ministry of Health. To estimate direct non-medical costs and annual gross earnings, estimates were obtained from a previous study that collected data using a direct non-medical costs patient questionnaire from a previous study on women diagnosed with breast cancer and receiving follow-up care at Mbabane Government chemotherapy unit (outpatient) in 2018 [22].

# Management of prostate cancer in Eswatini

In Eswatini, routine prostate cancer screening is only recommended for men above age 50 every after two years [23]. The referral pathway shown in Fig. 1, simplifies the treatment pathway which begins by a man presenting with symptoms or eligible for screening at outpatient. Patient will be referred to urologist for screening tests including PSA and digital rectal examination (DRE) [6, 23]. These tests are not confirmatory however, they



**Fig. 1** Simplified diagnosis and treatment pathway of patients diagnosed with prostate cancer. **PSA** Prostate specific antigen, **DRE** Digital rectal examination, **TURP** transurethral resection of prostate, **TURB** transurethral resection of bladder, **TRUS** Transrectal ultrasound, **LUTS** Lower Urinary Tract Symptoms

indicate changes in the prostate. Abnormal findings by either of the tests warrant further evaluation of patient and subsequent diagnostic test. These include biopsy (transrectal/perineal ultrasound guided biopsy (TRUS)). Patient with no cancer but presenting with symptom would receive management of lower urinary tract symptoms (LUTS). If cancer is confirmed further evaluation is conducted for cancer staging purposes in order to inform cancer management plan (metastasis screening). The evaluation includes radiology tests (bone scan, CT-scan and MRI pelvis). Staging is based on the tumor size (T) extent of lymph nodes involvement (N) and evidence of distant metastasis (M) [23, 24]. Depending on the risk score and prostate cancer stage, treatment include watchful waiting (cancer is monitored but not treated), surgery, radiation, chemotherapy and hormonal therapy (Androgen Deprivation Therapy) [23].

Most treatment modalities can be administered in various stages however for different intent [6, 23]. Radical prostatectomy, radiation and hormonal therapy can be applied for localised high risk prostate cancer (stage I and stage II) whilst for metastatic prostate cancer hormonal therapy will be first line in addition to radiotherapy, chemotherapy and hormonal therapy for palliation purposes. Radiation is not available in Eswatini and patient are referred to private hospitals in South Africa. Other surgical interventions for relieving symptoms such as

transurethral resection of prostate (TURP) or bladder (TURB) can be conducted locally.

We used expert opinion from Mbabane Government Hospital - Chemotherapy Unit, Mbabane clinic -private hospital and information from Phalala Fund to establish patient referral pathway. Phalala Fund is a government funded scheme established to fund provision of specialized health care services to people of Eswatini that could not afford payment of specialized care that is not available in country [25]. The Eswatini standardized cancer care guidelines were used to establish screening, diagnosis and treatment variables. Costs were estimated based on market price. Radiotherapy is currently not available in Eswatini. As such, patients who require radiation are managed in South Africa through Phalala Fund. Chemotherapy is available locally through a government chemotherapy unit and local private clinic. However, it was established that most patients were still receiving chemotherapy from South Africa.

# Costs

From a societal perspective, costs associated with prostate cancer were estimated to assess economic burden of prostate cancer in Eswatini. Direct medical costs were divided into recurrent and capital costs [19]. Recurrent costs included personnel, travel, consumables including medical supplies, administration, utilities and overheads.

Capital costs consistent mainly of equipment, building, vehicle and everything that have a useful life of more than one year. Costs for prostate cancer were determined based on the data source presented in Table 1. All costs were presented in US Dollars using 2018 average exchange rate (1 USD (\$) = 14.5 SZL).

# **Direct medical costs**

Direct costs in this study include resource utilization for diagnosis, treatment (surgery, chemotherapy, radiotherapy and androgen deprivation therapy) and follow-up care. To estimate the directs costs, we estimated average cost of each intervention from screening, staging and treatment and multiplied by the number of corresponding patients who received the intervention. The number of men diagnosed with prostate cancer were obtained from the national cancer registry [26]. All diagnosed cases were assumed to have undergone screening test using PSA. Screening and diagnosis costs were obtained from private hospital and market pricing. Treatment costs mainly radiation, chemotherapy and androgen deprivation therapy were received from Phalala fund based on South African private hospitals fees. In Eswatini, a majority of the management costs are borne by the Eswatini Government through Phalala fund.

As per the standardized cancer care guidelines, we assumed that all the men with confirmed prostate cancer in 2018 underwent screening and diagnosis tests, treatment and incurred other direct costs including transport and accommodation. Follow-up care costs was estimated for one year for those reported alive in 2018.

### **Direct non-medical costs**

Transport cost including return was estimated based on required patient follow-up visits based on the Eswatini Standardized Cancer Care Guidelines which state that follow-up visit should be every six months for the first two years and annually for up to five years following surgery [23]. Transport cost was estimated based on data from a previous study on breast cancer women receiving follow-up care at Mbabane Government Cancer Unit [22]. We assumed that all men completed treatment in 2018 had follow-up visits as per the Eswatini Standardized Cancer Guidelines. This study estimated one-year follow-up costs.

# **Indirect costs**

We estimated the monitory value of prostate cancer related productivity loss due to morbidity (patient sick

Table 1 Data variables and source for cost regarding screening, management and treatment of prostate cancer

| Data  | Data source  | Price source     |  |
|---|--|------------------|--|
| Estimated number of cases in 2018 = 90  | Swaziland National Cancer Unit, Eswatini Prostate cancer cases in 2018 | N/A              |  |
| Screening   |  |                  |  |
| Consultation fee  | Mbabane Clinic   | Private hospital |  |
| Prostate Specific Antigen (PSA)   | Eswatini Health Laboratory Services                                    | Private hospital |  |
| Digital rectal examination (DRE)  | Interview with expert  | Private hospital |  |
| Diagnosis   |  |                  |  |
| TRUS guided Biopsy  | Mbabane Clinic   | Private hospital |  |
| Computed Tomography (CT scan)   | Mbabane Clinic   | Private hospital |  |
| MRI scan  | Phalala fund   | Private hospital |  |
| X-ray   | Mbabane Clinic   | Private hospital |  |
| Bone scan   | Mbabane Clinic   | Private hospital |  |
| Intervention/Treatment  |  |                  |  |
| Watchful waiting (WW)   | Interview with expert  | Private hospital |  |
| Surgery   | Mbabane clinic   | Private hospital |  |
| Radiotherapy  | Phalala Fund based on SA hospitals fees                                | Private hospital |  |
| Chemotherapy  | Phalala Fund based on SA hospitals fees                                | Private hospital |  |
| Androgen deprivation  | Phalala Fund based on SA hospitals fees                                | Market price     |  |
| Hospitalization (local)   | Phalala Fund based on SA hospitals fees                                | Private hospital |  |
| Other direct costs  |  |                  |  |
| Transport and lodging costs in South Africa   | Phalala Fund based on SA hospitals fees                                | Market price     |  |
| Follow-up care (Year 1 following completion of treatment) Follow-up involves PSA testing, symptomology and clinical examination for metastatic cancer twice in a year | Based on reported prevalence   | Private hospital |  |

leave days incurred as a result of seeking health care) and pre-mature mortality).

The human capital method was used to estimate indirect costs related to productivity loss due to morbidity (sick leave as a result of seeking prostate cancer care) and pre-mature mortality [20]. We used average annual gross earnings computed from our previous study on breast cancer women receiving follow-up care in the chemotherapy unit, Mbabane Government hospital in Eswatini [22].

# Morbidity costs

We estimated the number of sick leave days for men diagnosed with prostate cancer who are in the labor participation ages (18-60 years). Using findings from a previously published study [27], we assumed sick leave for an average of 54 days per person. The sick leave days included days for staging, treatment and follow-up care. Using findings from a previous study on breast cancer conducted in Eswatini [22], we assumed 20 working days per month and a full-time working day of 8 h with estimated costs per workday (\$12) translating (\$1.5) per work hour [22].

# Mortality costs

To estimate the cost of lost productivity due to premature death related to prostate cancer, years of potential productive life lost (YPPLL) were calculated by subtracting age at death from the local retirement age of 60 years [28]. Prostate cancer age groups specific deaths were estimated assuming labor participation ages of Eswatini (18-60 years). We used full employment rate and annual average earnings obtained from a previous study. Average YPPLL was multiplied by average annual earnings. According to health economic recommendations, future costs were discounted at 3% and 5% [19, 29]. The number of prostate cancer related deaths was obtained from Eswatini Cancer Registry. In 2018, there were 31 prostate cancer related mortality with 4 that occurred within the labor participating ages of Eswatini (18-60) years [28].

# Cancer mortality and years of potential productive life lost (YPLL)

The number of prostate cancer related deaths was obtained from Eswatini Cancer Registry from which the years of productive life lost was calculated. In 2018, there were 31 prostate cancer related mortality out of which 4 occurred within the labor participating ages of Eswatini (18-60) years [28].

# **Estimation of annual costs**

We computed the aggregate total costs of screening, diagnosis and treatment of prostate cancer in 2018 as below:

Cost of disease = 
$$\sum \begin{pmatrix} Direct \ medical \ \cos ts \\ + \\ Direct \ non - medical \ \cos ts \\ Direct \ \cos ts \end{pmatrix} + \begin{pmatrix} Morbidity \\ + \\ Mortality \\ Indirect \ \cos ts \end{pmatrix}$$

Direct medical costs = Consisting of direct non-medical costs and direct medical costs.

Indirect costs = Consisting of morbidity costs and mortality costs (Patient time lost as a result of the condition and costs associated with premature mortality as a result).

All costs were reported in 2018 US dollars (\$1=SZL14.5).

# Sensitivity analysis

Sensitivity analysis was performed using  $\pm$  25% to account for the cost of follow-up prevalent cancer cases and to account for unrecorded cases by the facilities.

# **Results**

# **Directs costs**

In 2018, there were 90 prostate cancer cases of which 89% aged 60 years an above. The average age was 73 years. Table 2, shows unit average costs for treating prostate cancer cases including other direct costs such as transportation and accommodation.

Cost distribution by disease stage is shown in Table 3. Following the Eswatini Standard Cancer Care Guidelines we assumed that all confirmed cases underwent similar screening, diagnosis and treatment pathway shown in Table 2, and simplified referral pathway shown in Fig. 1. The average costs for the different pathway including treatment intervention differed with the prostate cancer stage. Radical prostate comy was more frequent with early stages of prostate cancer whilst interventions like chemotherapy were common with prostate cancer stages III and IV. Table 3 shows the prostate cancer costs distribution by stage.

Radiation is not available in Eswatini and patients are referred to private hospitals in South Africa. On average, radiotherapy treatment is administered for a period of 5-weeks [25]. The estimated unit costs for radiotherapy was \$16,648 whilst chemotherapy was \$7,498. In addition to treatment costs, all patients referred for radiotherapy also incurred other direct costs including transport, lodging and allowance for accompanying staff (nurse and driver) at a unit costs \$5,959, Table 3.

Table 2 Costs for screening, diagnosis and treatment of prostate cancer

| Parameter   | Variables included in the cost  | Average (2018)<br>USD |
|---|---|-----------------------|
| Screening   | Consultation fee  | 41                    |
|   | Prostate Specific Antigen (PSA)   | 16                    |
|   | Digital rectal examination (DRE)  | 32                    |
| Diagnosis   |   |                       |
| Pathology   | TRUS guided Biopsy  | 147                   |
| Radiology   | Computed Tomography (CT scan)- to rule out chest, abdomen and pelvis metastasis                                       | 862                   |
|   | Magnetic resonance imaging scan (MRI)   | 1,034                 |
|   | X-ray to rule out effusion  | 28                    |
|   | Bone scan in locally advanced prostate cancer   | 607                   |
|   | Ultrasound scan   | 103                   |
| Treatment   |   |                       |
| Watchful Waiting (WW) cost include PSA test every three months and follow up consultation fee   | PSA plus follow-up consultation fee   | 58                    |
| Surgery   | Radical prostatectomy   | 5,726                 |
|   | Orchiectomy   | 5,726                 |
| Radiotherapy  | Administered at 5 function of a 5-week period   | 16,647                |
| Chemotherapy (Brachytherapy or external beam radiation  | Administered in 5 cycles over 5-week period   | 7,498                 |
| Androgen Deprivation Therapy (ADT)  | Mostly Zoladex o.8 mg intramuscular (IM) for every 3 months   | 1,268                 |
| Symptoms relieving procedures (TURP/TURB)   | Transurethral resection of the prostate or bladder (TURP/TURB)  | 5,872                 |
| Other direct costs  |   |                       |
| Hospitalization costs (local)   | Admitted for symptoms management procedures including transurethral resection of the prostate (TURP) and orchidectomy | 5,872                 |
| Hospital admission in step down facility for late stage treatment (In South Africa)   | Patient who require close monitoring following radiotherapy or surgery in hospital outside Eswatini                   | 1,206                 |
| Transport and lodging cost in South Africa  | All patients who received treatment in South Africa   | 5,959                 |
| Follow-up care (Year 1 following completing treatment) Follow-up is done using PSA testing, symptomology and clinical examination for metastatic cancer twice in a year | Follow up consultation, PSA tests.  | 94                    |
| Total   |   | 58,796                |

# Direct non-medical costs

Using estimate from a previous study [22], the average transport cost per follow-up visit including return was \$11 (inter quartile range (IQR)\$4-46). On average, post treatment follow-up visits should be every 6 months resulting to four visits in a year including return. We assumed that all patients visited the hospital in the company of a relative. The total average transport costs including return was estimated at \$5,029 (between \$3,771 and 6,287 estimated with lower and upper bounds).

# Indirect costs

Productive loss due to sick leave as a result of patient seeking health care for prostate cancer was estimated at \$58,320, Table 4. Out of the 90 patients diagnosed with prostate cancer, there were 13 men within the labor participating ages which were assumed to be on average sick leave of 54 days per person excluding short term

sick leave of 14 days that is usually covered by employers. A total of 31 men died of prostate cancer in 2018 out of which 4 were less than 60 years. Costs due to prostate cancer premature mortality was estimated at \$113,760, Table 5.

# **Total annual costs**

The total annual costs for prostate cancer was estimated at \$ 6.2 million (between \$4.7 million and 7.8 million estimated with lower and upper bounds), Table 6. Fourth 4% (40) of the cases were diagnoses with stage IV whilst only 11% (10) were diagnosed with stages I. Management of prostate cancer stages III and IV formed the greatest share of the costs for prostate cancer contributing about \$1.2 and 2.1 million respectively. The total costs of stages I and II was estimated at \$0.5 and \$0.8 million. Transport and accommodation costs (cost incurred by those transferred to South Africa) were highest under other direct

Table 3 Costs for staging, management, and treatment of Prostate cancer stage I-IV

| Staging and treatment variables   | Unit cost (\$) | I (T1) | II (T2) | III (T3) | 1 V (T4) |
|---|----------------|--------|---------|----------|----------|
| Consultation for assessment   | 41             | 41     | 41      | 41       | 41       |
| Screening and diagnosis   |                |        |         |          |          |
| Prostate Specific Antigen (PSA)   | 16             | 48     | 48      | 48       | 48       |
| Digital rectal examination (DRE)  |                |        |         |          |          |
| TRUS guided Biopsy  | 147            | 147    | 147     | 147      | 147      |
| MRI scan  | 1,034          | 1,034  | 1,034   | 1,034    | 1,034    |
| Chest x-ray   | 28             | 27.6   | 27.6    | 27.6     | 27.6     |
| Bone scan   | 607            | 607    | 607     | 607      | 607      |
| Ultrasound  | 103            | 103    | 103     | 103      | 103      |
| CT scan abdomen   | 862            | 862    | 862     | 862      | 862      |
| Treatment (Prostate Cancer prevalence in 2018=91 patient)                                       |                |        |         |          |          |
| Watchful waiting (WW). Costs include PSA test every three months and follow up consultation fee | 58             | 58     | 0       | 0        | 0        |
| Radical prostatectomy   | 5,726          | 5,726  | 5,726   | 0        | 0        |
| Orchiectomy (surgical castration)   | 5,726          | 5,726  | 5,726   | 5,726    | 5,726    |
| Radiotherapy  | 16,648         | 16,648 | 16,648  | 16,648   | 16,648   |
| Chemotherapy  | 7,498          | 0      | 0       | 7,498    | 7,498    |
| Symptoms relieving procedures (TURP/TURB)   | 5,872          | 0      | 0       | 5,872    | 5,872    |
| Other supportive drugs: Pain killers  | 60             | 60     | 60      | 60       | 60       |
| Hormonal therapy (ADT) Zoladex 0.8 mg injectables   | 1,268          | 1,268  | 1,268   | 1,268    | 1,268    |
| Other costs   |                |        |         |          |          |
| Hospitalization costs (local)   | 5,872          | 5,872  | 5,872   | 5,872    | 5,872    |
| Hospital admission in step down facility for late stage treatment                               | 1,206          | 1,206  | 1,206   | 1,206    | 1,206    |
| Transport and lodging cost (in RSA)   | 5,959          | 5,959  | 5,959   | 5,959    | 5,959    |
| Follow-up care (Year 1 following completing treatment)  | 94             | 94     | 94      | 94       | 94       |
| Total   | 58,824         | 45,486 | 45,428  | 53,072   | 53,072   |

**Table 4** Costs due to sick leave days associated with prostate cancer costs

|               | Numbers of<br>sick leave<br>days | Number<br>of patients<br>alive in<br>2018 | Cost per<br>workday (\$) | Total<br>productivity<br>loss due to<br>costs due<br>to prostate<br>cancer in<br>2018 (\$) for<br>all patient |
|---------------|----------------------------------|---|--------------------------|---|
| Total patient | 54                               | 58  | 12                       | \$37,584  |

costs contributing about \$0.5million. In 2018, there were 31 prostate cancer related deaths with only 4 occurred within the labor participating ages of Eswatini (18-60) years. The total year of productive life lost (YPPL) was 221 years. Indirect costs were estimated at \$0.24 million and a majority (96%, \$0.2 million) were productive loss from premature mortality, Table 6.

# **Discussion**

The current study assessed the costs associated with prostate cancer in Eswatini, that is, screening, diagnosis, treatment and follow-up care. The study considered direct costs including follow-up care costs within one year of diagnosis. To our knowledge this is the first study to estimate the economic burden of prostate cancer in Eswatini. The estimated annual prostate cancer burden was \$ 6.1 million in 2018. About 89% of the patient aged 60 years and above. Given the Eswatini Standardized Cancer Care and Guidelines [21], we assumed that all patients diagnosed in 2018 underwent the screening and diagnostic procedures. Treatment costs varied by cancer stage reflecting the utilization of treatment modalities per stage hence high costs observed in stages III (\$1.2million) and IV (2.1million) versus Stage I and II with \$0.5 and \$0.8 million respectively. The findings indicate that managing advanced stages of the disease increases health care costs.

The study findings were in accordance with findings from other studies. A study assessing health care costs associated with prostate cancer in Canada reported

**Table 5** Mortality for prostate cancer

| Mortality cost for Prostate cancer |  |  |                             |   |   |  |
|------------------------------------|--|--|-----------------------------|---|---|--|
| Age groups                         | Lost YPPLL (Average YPPLL<br>for 1 patient in each age<br>group) | Number of premature deaths before age 60 | Average<br>annual<br>income | Mortality cost (\$) with 3%<br>Discount rate multiplied<br>with the number of<br>patients in this age group | Mortality cost (\$) with 5%<br>Discount rate multiplied<br>with the number of<br>patients in each age group |  |
| 46-51                              | 49   | 1  | 2690                        | 29,632  | 10,676  |  |
| 52-56                              | 54   | 3  | 2690                        | 84,128  | 27,311  |  |
| Totals                             | 103  | 4  | 2690                        | 113,760   | 37,987  |  |
| YPPLL                              | 211  |  |                             |   |   |  |
| Average annual g                   | gross income = \$2,690   |  |                             |   |   |  |

increasing costs per stage I (\$1,297), II (\$3,289), III (\$1,495), IV (\$5,629) and V (\$16,020) [30]. Similarly, a study conducted in Iran concluded that health care costs for metastatic stages were the highest compared to treatment costs for localized prostate cancer [11]. More studies had similar conclusions [31, 32]. Slightly different findings were from the United State of America who reported high treatment costs for initial diagnosis and metastatic phase with radical prostatectomy being the main cost driver [33]. Whilst in this study we found lesser cost with early stage cancer, however, both studies observed increasing costs with advanced cancer stages. Also, the differences could be partly explained by the men (20%) diagnosed with early stages of prostate cancer in our study. A systematic review of registry-based studies assessing economic burden of prostate cancer in Europe found that cost distribution across prostate cancer stages varied across countries [34]. This can be attributed to differences in prostate cancer detection and country specific management practice [34]. The authors also acknowledged the difference in methodologies applied in the studies as possible explanation to the varying outcome

There seems to be lack of global consensus on prevention strategies particularly age of screening. The United State Preventive Service Task Force (USPSTF) recommend against routine screening for men 70 years and older for prostate cancer particularly using prostate specific antigen screening [35]. The Eswatini Standardized Cancer Care and Guidelines also discourages routine prostate cancer screening with an exception for men 50 years and above or symptomatic [23]. Other studies argue that increased screening lead to increased detection of low-grade cancers resulting to patient with indolent tumors receiving aggressive treatment [36].

In LICs such as Eswatini, the challenge is likely to be on a different direction than over diagnosing and consequently overt treatment. Lack of screening and comprehensive treatment remains the greatest challenge for most LMICs and LICs. Eswatini is not different from other low middle income countries from whom late diagnosis coupled with limited treatment options remains a challenge. In Eswatini, in 2018, more than 80% of the patients were diagnosed with advanced cancer (stages III and IV), yet major treatment is not available in country. These include radiotherapy and androgen deprivation therapy (ADT). Accessing care outside the country comes with additional costs, mainly accommodation, transportation and meals for patients referred to South Africa.

Lack of specialized and costly care have been reported in other countries particularly in Africa and mortality from prostate cancer is the highest in these countries and there is lack of cancer treatment guidelines [4, 37].

There is an urgent need to strengthen health systems enablers [38]. These include investments in the establishment of local cancer treatment centers, optimizing health workforce competencies throughout the continuum of care and ensuring availability of medical products and diagnostics technologies to facilitate local diagnosis, staging and management.

Despite the evidence that prostate cancer is a major public health challenge, literature on the economic burden of prostate cancer is however limited and severely so in low income countries particularly in the sub-Saharan region. Findings from a systematic review on the costs of prostate cancer studies indicated a need not only for harmonized methodologies but also to expand research in this field [39]. Similarly, another systematic literature review of registry-based studies reached similar conclusion on the need for further research in cost of illness studies focusing on prostate cancer [40].

In the study we assessed indirect costs by estimating the costs associated with unpaid sick leave days and productive loss due to premature mortality from prostate cancer. Of the total costs, indirect costs accounted for 4.2% (\$0.24 million). Comparing these findings to previous cost analysis studies for prostate cancer, most of the

**Table 6** Total Annual costs estimation for Prostate cancer (direct and indirect costs)

|   | Prevalence 2018 | Cost per item (\$)  | Base case cost (\$) | Range (\$)    |              |
|---|-----------------|---------------------|---------------------|---------------|--------------|
| Parameter   | Number          | Average cost (2018) | Base costs (2018)   | (Lower (-25%) | Higher (+25) |
| Direct costs (Health care costs) consultation fee                 | 90              | 41                  | 3690                | 2768          | 4613         |
| Screening and diagnosis   |                 |                     |                     |               |              |
| Prostate Specific Antigen (PSA)                                   | 90              | 16                  | 1448                | 1086          | 1810         |
| Digital rectal examination (DRE)                                  | 8               | 0                   | 0                   | 0             | 0            |
| TRUS guided Biopsy  | 90              | 147                 | 13,213              | 9910          | 16,516       |
| MRI scan  | 90              | 1034                | 93,060              | 69,795        | 116,325      |
| Chest x-ray   | 90              | 28                  | 2484                | 1863          | 3105         |
| Bone scan   | 90              | 607                 | 54,630              | 40,973        | 68,288       |
| Ultra sound   | 90              | 103                 | 9270                | 6953          | 11,588       |
| CT scan abdomen   | 90              | 862                 | 77,580              | 58,185        | 96,975       |
| Treatment   |                 |                     |                     |               |              |
| Stage I   | 10              | 45,486              | 454,861             | 341,146       | 568,577      |
| Stage II  | 17              | 45,428              | 772,278             | 579,209       | 965,348      |
| Stage III   | 23              | 53,072              | 1,220,659           | 915,494       | 1,525,824    |
| Stage IV  | 40              | 53,072              | 2,122,886           | 1,592,164     | 2,653,607    |
| Other direct costs  |                 |                     |                     |               |              |
| Hospitalization costs (local)                                     | 90              | 210                 | 18,900              | 14,175        | 23,625       |
| Hospital admission in step down facility for late stage treatment | 90              | 1206                | 108,540             | 81,405        | 135,675      |
| Transport and lodging cost (in RSA)                               | 90              | 5959                | 536,310             | 402,233       | 670,388      |
| Follow-up care (Year 1 following completing treatment)            | 60              | 2662                | 159,720             | 119,790       | 199,650      |
| Total direct  |                 | 209,892             | 5,645,839           | 4,234,379     | 7,057,299    |
| Direct non-medical cost   |                 |                     |                     |               |              |
| Transport costs for follow-up visits ,patient                     | 59              | 2513                | 148,267             | 111,200       | 185,334      |
| Transport costs for follow-up visits, accompanying relative       | 59              | 2513                | 148,267             | 111,200       | 185,334      |
| Total Direct non-medical costs                                    |                 | 5026                | 296,534             | 222,401       | 370,668      |
| Indirect costs  |                 |                     |                     |               |              |
| Morbidity costs due to sick leave                                 | 13              | 648                 | 8424                | 6318          | 10,530       |
| Premature mortality costs   | 4               | 57,675              | 230,700             | 173,025       | 288,375      |
| Total indirect costs  |                 | 58,323              | 239,124             | 179,343       | 298,905      |
| Total   |                 | 268,215             | 6,181,497           | 4,636,123     | 7,726,871    |

studies did not consider assessing indirect costs, however a study from Sweden reported low proportion of productivity loss associated with prostater cancer [9]. Further comparison of the findings with studies from other cancer types conducted in Eswatini [22, 41], the indirect costs from this study accounted for a lesser share of the total cost. This could partly be explained by the fact that most participants (89%) were above the labor participating ages (18-60 years) and few deaths occurred below age 60 years. A similar pattern was observed in Sweden, again the finding were attributed to low number of prostate cancer cases and deaths among labor participation groups [9].

The key strength of our study was that this is the first study to estimate cost associated with prostate cancer in Eswatini. The study considered both direct and indirect costs of prostate cancer. Our study has notable findings that has implications on health care systems strengthening and resources allocation in Eswatini. Our study present description of resource utilization and associated health care costs in managing prostate cancer in Eswatini.

An important limitation is the absence of index cost in Eswatini. We considered private and market prices for best possible price estimates.

The estimates presented were based on available data however, estimates could be conservative due to several reasons, First, due to limited data availability we used information from literature and interview with experts for some treatment variables, as such, some information can be subject to context and preferences. Secondly, we only considered costs in the first year of diagnosis yet cost for follow-up care can be even beyond five years [6, 42]. Lastly, we employed human capital approach to estimate the costs related to productivity loss associated with prostate cancer. Whilst this is a commonly applied approach, it is mostly criticized for excluding individuals above the labor participation age group yet there is argument that some of those people can still be involved in labor activities that gives meaningful income. Another author argues that this has severe implication when valuing productivity loss for prostate cancer given that a majority of the patients are diagnosed after they have past the retirement age.

### **Conclusions**

The findings of the study indicated that costs attributed to prostate cancer were substantial and they are a public health concern. The findings were consistent with those of other countries, a majority of which were conducted in developed countries. The study demonstrated the interventions and associated costs. Radiotherapy was the most expensive treatment intervention in Eswatini, yet other studies cited surgery related intervention as the major costs driver. This is a reasonable finding in the context of Eswatini given that radiotherapy treatment is not available locally, patients are referred to private hospitals outside the country. The findings point areas for policy makers to perform cost containment regarding therapeutic procedures for prostate cancer. Also, the study findings demonstrate that prostate cancer costs are likely to increase in future and there is a need for strengthening adherence to the Eswatini Standardized Cancer Care and Guidelines in order to ensure that resources are invested to diagnosing the most at risk groups.

# Abbreviations

ADT: Androgen Deprivation Therapy; DRE: Digital Rectal Examination; PSA: Prostate Specific Antigen test; TRUS: Transrectal ultrasound; LUTS: Low Urinary Symptom; HICs: High Income Countries; LICs: Low Income Countries; LMICs: Low Middle Income Countries; SDGs: Sustainable Development Goals.

# **Supplementary information**

The online version contains supplementary material available at https://doi.org/10.1186/s12913-022-07817-6.

Additional file 1. Direct non-medical costs patient questionnaires.pdf

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### Authors' contributions

Conceptualization: CN, Data curation: CN TG EO, Formal analysis: CN TG EO, Funding acquisition: CN, Investigation: CN, Methodology: CN TG EO, Project administration: CN, Supervision: TG EO, Validation: TG EO, Writing – original draft: CN, Writing – review & editing: CN TG EO. The author(s) read and approved the final manuscript.

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### Availability of data and materials

All data generated or analyzed during this study are included in this published article. See study tables and figures.

# **Declarations**

### Ethics approval and consent to participate

The study was approved by the National Health Research Review Board (NHRRB) of Eswatini (FWA 00026661/IRB 00011253) and the Biomedical Research Ethics Committee of the University of KwaZulu-Natal (BE 059/19). Ethics committees approved written informed consent, which was obtained from all the participants prior to participation to the study. The study was implemented in accordance with the approved protocol and ethics principles quiding human participation in research.

# Consent for publication

Not applicable.

### Competing interests

The authors declare that they have no competing interests.

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